NDA 19-012/S-017 AUG 25 2000

McNeil Consumer Healthcare
Attention: Janet A. Uetz
Associate Director, Regulatory Affairs
7050 Camp Hill Road
Fort Washington, PA 19034-2299

Dear Ms. Uetz:

Please refer to your supplemental new drug application dated March 15, 1999, received March 16, 1999, submitted under section 505(b) of the Federal Food, Drug, and Cosmetic Act for Motrin lB (ibuprofen) Tablets and Caplets, 200 mg.

This "Changes Being Effected" supplemental new drug application provides for revised labeling to implement the allergy alert statements required by our September 15, 1998 letter, and the alcohol warning required by the final rule published on October 23, 1998 (63 FR 56789).

We have completed the review of this supplemental new drug application and have concluded that adequate information has been presented to demonstrate that the drug product is safe and effective for use as recommended in the final printed labeling submitted on March 15, 1999. Accordingly, the supplemental new drug application is approved effective on the date of this letter.

We request that the following revisions in the labeling for this drug product be implemented within 180 days or at the next printing, whichever comes first:

1. Only the first letter in the header "Alcohol warning" should be capitalized, all other letters should be in lower case.

2. Only the first letter in the header "Allergy alert" should be capitalized, all other letters should be in lower case.

3. A bullet point (e.g., solid circle or solid square) should precede the statements under the additional warnings listed below. The period should be deleted after the first bulleted statement under the subheading "Do not use" as specified below:

   Do not use
   • if you have ever had an allergic reaction to any other pain reliever/fever reducer
   Stop use and ask a doctor if
   • an allergic reactions occurs. Seek medical help right away.

4. The storage statement should read "store at 20- 25°C (68 - 77°F)."

5. The phrase "No prescription needed" should be deleted since the product has been available as an OTC product for greater than 6 months.

Furthermore, we note that the labeling was not submitted in Drug Facts format consistent with the requirements of the March 17, 1999 FEDERAL REGISTER document "Over-the-Counter Human Drugs; Labeling Requirements; Final Rule" (64 FR 13254) (OTC labeling final rule), which has been incorporated into the regulations at 21 CFR 201.66. We remind you that the labeling of your product must be revised to reflect the Drug Facts format within the timeframes specified in the OTC labeling final rule.

If a letter communicating important information about this drug product (i.e., a "Dear Health Care Practitioner" letter) is issued to physicians and others responsible for patient care, we request that you submit a copy of the letter to this NDA and a copy to the following address:

MED WATCH, HF-2
FDA
5600 Fishers Lane
Rockville, MD 20857.

We remind you that you must comply with the requirements for an approved NDA set forth under 21 CFR 314.80 and 314.81.

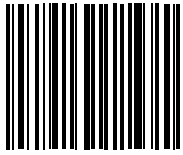
CONSUMER LABELING LEAFLET FOR MOTRIN® IB. PLEASE SAVE THIS FOR FUTURE USE.

Only selected information is contained on the bottle label. Therefore, you should keep this sheet for future reference.

Motrin®

IB IBUPROFEN TABLETS USP

Pain Reliever/ Fever Reducer



WARNINGS:

Allergy Alert: ibuprofen may cause a severe allergic reaction which may include:

- **hives**
- **facial swelling**
- **asthma (wheezing)**
- **shock**

Do not use if you have ever had an allergic reaction to any other pain reliever/fever reducer.

Stop use and ask a doctor if an allergic reaction occurs. Seek medical help right away.

INDICATIONS: For the temporary relief of headache, muscular aches, the minor pain of arthritis, toothache, backache, minor aches and pains associated with the common cold, the pain of menstrual cramps, and for reduction of fever.

DIRECTIONS: Adults: Take 1 tablet (or caplet*) every 4 to 6 hours while symptoms persist. If pain or fever does not respond to 1 tablet, 2 tablets may be used, but do not exceed 6 tablets in 24 hours, unless directed by a doctor. The smallest effective dose should be used. Take with food or milk if occasional and mild heartburn, upset stomach or stomach pain occurs with use. Consult a doctor if these symptoms are more than mild or if they persist.

Children: Do not give this product to children under 12 except under the advice and supervision of a doctor.

ALCOHOL WARNING: If you consume 3 or more alcoholic drinks every day, ask your doctor whether you should take ibuprofen or other pain relievers/fever reducers. Ibuprofen may cause stomach bleeding.

WARNINGS: Do not take for pain for more than 10 days or for fever for more than 3 days unless directed by a doctor. If pain or fever persists or gets worse, if new symptoms occur, or if the painful area is red or swollen, consult a doctor. These could be signs of serious illness. If you are under a doctor's care for any serious condition, consult a doctor before taking this product. As with aspirin and acetaminophen, if you have any condition which requires you to take prescription drugs, or if you have had any problems or serious side effects from taking any non-prescription pain reliever, do not take MOTRIN® IB without first

(continued on back)

discussing it with your doctor. If you experience any symptoms which are unusual or seem unrelated to the condition for which you took ibuprofen, consult a doctor before taking any more of it. Although ibuprofen is indicated for the same conditions as aspirin and acetaminophen, it should not be taken with them except under a doctor's direction. Do not combine this product with any other ibuprofen-containing product.

Do not exceed recommended dose. Keep this and all drugs out of the reach of children. In case of accidental overdose, seek professional assistance or contact a poison control center immediately. As with any drug, if you are pregnant or nursing a baby, seek the advice of a health professional before using this product. IT IS ESPECIALLY IMPORTANT NOT TO USE IBUPROFEN DURING THE LAST 3 MONTHS OF PREGNANCY UNLESS SPECIFICALLY DIRECTED TO DO SO BY A DOCTOR BECAUSE IT MAY CAUSE PROBLEMS IN THE UNBORN CHILD OR COMPLICATIONS DURING DELIVERY.

Store at room temperature. Avoid excessive heat 40°C (104°F).

Active Ingredient: **Each tablet (or caplet*) contains ibuprofen 200 mg.**

Inactive Ingredients: **Carnauba wax, cornstarch, hydroxypropyl methylcellulose, iron oxide black, pregelatinized starch, propylene glycol, silicon dioxide, stearic acid, titanium dioxide.**

*Capsule-Shaped Tablets

815490102
691346

McNeil

Distributed by
McNeil Consumer Healthcare
DIVISION OF MCNEIL-PPC, INC.
FORT WASHINGTON, PA 19034 USA

To Open:

1. Hold bottle firmly with one hand.
2. While pushing down with other hand, turn cap counter-clockwise.

To Close:

1. Place cap on bottle.
2. Turn cap clockwise until TIGHTLY in place.


See New Label
No Prescription Needed
Motrin®
IB IBUPROFEN TABLETS USP
Pain Reliever/ Fever Reducer
100 COATED 200 mg each IBUPROFEN CAPLETS
Caplets*
*Capsule-Shaped Tablets
Do not use if neck wrap or foil inner seal imprinted "Safety Sealed" is broken or missing.
McNeil
Distributed by
McNeil Consumer Healthcare
DIVISION OF MCNEIL-PPC, INC.
FORT WASHINGTON, PA 19034 USA
©MCN-PPC, Inc. '98
3 0045-0481-01 6
773348
NOT VALID FOR PROOF-OF-PURCHASE
0815312207
Motrin IB IBUPROFEN TABLETS USP Pain Reliever/Fever Reducer
WARNINGS:
Allergy Alert: ibuprofen may cause a severe allergic reaction which may include:
• hives • facial swelling
• asthma (wheezing) • shock
Do not use if you have ever had an allergic reaction to any other pain reliever/fever reducer.
Stop use and ask a doctor if an allergic reaction occurs. Seek medical help right away.
INDICATIONS: For the temporary relief of headache, muscular aches, the minor pain of arthritis, toothache, backache, minor aches and pains associated with the common cold, the pain of menstrual cramps, and for reduction of fever.
DIRECTIONS: Adults: Take 1 caplet every 4 to 6 hours while symptoms persist. If pain or fever does not respond to 1 caplet, 2 caplets may be used, but do not exceed 6 caplets in 24 hours, unless directed by a doctor. The smallest effective dose should be used. Take with food or milk if occasional and mild heartburn, upset stomach or stomach pain occurs with use. Consult a doctor if these symptoms are more than mild or if they persist.
Children: Do not give this product to children under 12 except under the advice and supervision of a doctor.
Active Ingredient: Each caplet contains ibuprofen 200 mg.
Inactive Ingredients: Carnauba wax, cornstarch, hydroxypropyl methylcellulose, iron oxide black, pregelatinized starch, propylene glycol, silicon dioxide, stearic acid, titanium dioxide.
See end panel for expiration date.
Store at room temperature. Avoid excessive heat 40°C (104°F).
815 312 207
ALCOHOL WARNING: If you consume 3 or more alcoholic drinks every day, ask your doctor whether you should take ibuprofen or other pain relievers/fever reducers. Ibuprofen may cause stomach bleeding.
WARNINGS: Do not take for pain for more than 10 days or for fever for more than 3 days unless directed by a doctor. If pain or fever persists or gets worse, if new symptoms occur, or if the painful area is red or swollen, consult a doctor. These could be signs of serious illness. If you are under a doctor's care for any serious condition, consult a doctor before taking this product. As with aspirin and acetaminophen, if you have any condition which requires you to take prescription drugs, or if you have had any problems or serious side effects from taking any non-prescription pain reliever, do not take MOTRIN® IB without first discussing it with your doctor. If you experience any symptoms which are unusual or seem unrelated to the condition for which you took ibuprofen, consult a doctor before taking any more of it. Although ibuprofen is indicated for the same conditions as aspirin and acetaminophen, it should not be taken with them except under a doctor's direction. Do not combine this product with any other ibuprofen-containing product.
Do not exceed recommended dose. Keep this and all drugs out of the reach of children. In case of accidental overdose, seek professional assistance or contact a poison control center immediately.
As with any drug, if you are pregnant or nursing a baby, seek the advice of a health professional before using this product. IT IS ESPECIALLY IMPORTANT NOT TO USE IBUPROFEN DURING THE LAST 3 MONTHS OF PREGNANCY UNLESS SPECIFICALLY DIRECTED TO DO SO BY A DOCTOR BECAUSE IT MAY CAUSE PROBLEMS IN THE UNBORN CHILD OR COMPLICATIONS DURING DELIVERY.
Lot
Exp
Do not use if neck wrap or foil inner seal imprinted "Safety Sealed" is broken or missing.
WARNINGS:
Allergy Alert: ibuprofen may cause a severe allergic reaction which may include:
• hives • facial swelling
• asthma (wheezing) • shock
Do not use if you have ever had an allergic reaction to any other pain reliever/fever reducer.
Stop use and ask a doctor if an allergic reaction occurs. Seek medical help right away.
INDICATIONS: For the temporary relief of headache, muscular aches, the minor pain of arthritis, toothache, backache, minor aches and pains associated with the common cold, the pain of menstrual cramps, and for reduction of fever.
Motrin® IB IBUPROFEN TABLETS USP Pain Reliever/ Fever Reducer
100 COATED 200 mg each IBUPROFEN CAPLETS*
*Capsule-Shaped Tablets
See consumer labeling leaflet for complete usage information.
DIRECTIONS: Adults: Take 1 caplet every 4 to 6 hours while symptoms persist. If pain or fever does not respond to 1 caplet, 2 caplets may be used, but do not exceed 6 caplets in 24 hours, unless directed by a doctor. The smallest effective dose should be used. Take with food or milk if occasional and mild heartburn, upset stomach or stomach pain occurs with use. Consult a doctor if these symptoms are more than mild or if they persist.
Children: Do not give this product to children under 12 except under the advice and supervision of a doctor.
Active Ingredient: Each caplet contains ibuprofen 200 mg.
Store at room temperature. Avoid excessive heat 40°C (104°F).
814 492 309
©MCN-PPC, Inc. '98
DISTRIBUTED BY:
MCNEIL-PPC, INC.,
FORT WASHINGTON, PA 19034 USA